

Calcified brain metastatic adenocarcinoma: A case report and review of the literature

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Abstract

Introduction: Calcification in brain metastasis occurs rarely so it is reported in approximately 1% of surgical and 6.6% of autopsy specimens. Here we report a new case of brain metastasis with calcification.

Case presentation: A 44-year-old woman presented with a generalized tonic-clonic seizure with no neurological deficit on physical examination. Brain imaging demonstrated a hyperdense lesion on computed tomography scan and hyposignal and rim enhancement on T1, T2 and T1 with gadolinium injection sequence images in the right parieto-occipital lobe. Intraoperatively, there was a well-defined solid homogenous calcified mass within brain parenchyma. The lesion that resembled a meningioma was totally resected. The histopathological examination revealed metastatic adenocarcinoma.

Conclusion: Metastatic brain lesions should be in the differential diagnosis of a solitary calcified brain mass, although it occurs rarely. It is important to differentiate it radiologically from intralesional haemorrhage.

Keywords

Calcified brain mass, metastatic adenocarcinoma, colon cancer

Introduction

Calcification is a physiological process in the brain and also in the choroid plexus and pineal gland. As a pathological process, it occurs in vascular, neoplastic, inflammatory and metabolic diseases.¹ Brain tumours such as meningioma and oligodendroglioma, infectious processes such as tuberculosis, fungal and cysticercosis and metabolic diseases such as hyperparathyroidism are the most common clinical situations associated with an intracranial calcified mass.² Calcification in brain metastasis occurs rarely so it is reported in approximately 1.1% of surgical and 6.6% of autopsy specimens.³ Here we report a new case of brain metastasis with calcification.

Case presentation

A 44-year-old woman presented with a generalized tonic-clonic seizure 3 days before the admission. Physical examination showed no neurological deficit. Brain computed tomography (CT) scan demonstrated a hyperdense lesion in the right parieto-occipital lobe (Figure 1). Brain magnetic resonance imaging (MRI) demonstrated a well-defined lesion in the right parieto-occipital lobe that was hyposignal on T1 and T2 associated with rim enhancement on T1 with gadolinium injection sequence images (Figure 2). There was moderate vasogenic oedema around the lesion. Laboratory tests and chest X-ray were normal.

After right parietooccipital craniotomy, opening the dura by the transcortical approach, there was a well-defined solid homogenous calcified mass within the brain parenchyma. The lesion that resembled a meningioma was totally resected. The histopathological examination revealed metastatic adenocarcinoma, most probably from the colon region. The patient was referred to an oncologist for further work-up and treatment. After scalp wound healing, the patient received adjuvant cranial radiotherapy for local control of the disease. After a thorough work-up, the large intestine tumour (caecal adenocarcinoma) was affirmed, and she underwent systemic chemotherapy for treatment. At the last follow-up 4 months after surgery, she had completed the adjuvant radiotherapy and was receiving systemic chemotherapy, she had no neurological signs and symptoms and no evidence of recurrence on brain imaging.

Discussion

Metastatic brain lesions can be solitary or multiple. On CT scan, they can be hypo, iso or hyperdense

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compared to the adjacent brain parenchyma. However, in the setting of metastatic brain melanoma and intralésional haemorrhage, CT scan demonstrated a hyperdense lesion.⁴

Haemorrhage into the metastasis can make the diagnosis of calcified mass challenging. Some attention helps to differentiate them from each other. First, attenuation values for haemorrhage are usually 35–55 Hounsfield units, but two studies demonstrated Hounsfield units of 50–105 and 86 for calcification within metastasis.⁵ Second, it is more likely that punctuate and curvilinear hyperdensity is metastasis, but differentiation in the setting of amorphous hyperdensity is difficult. Third, calcification is usually unchanged or progresses very slowly during follow-up, but haemorrhage-induced hyperdensity resolves over time.⁵

Brain metastasis does not generally calcify. When a metastasis shows calcification other diagnoses should be considered.⁴

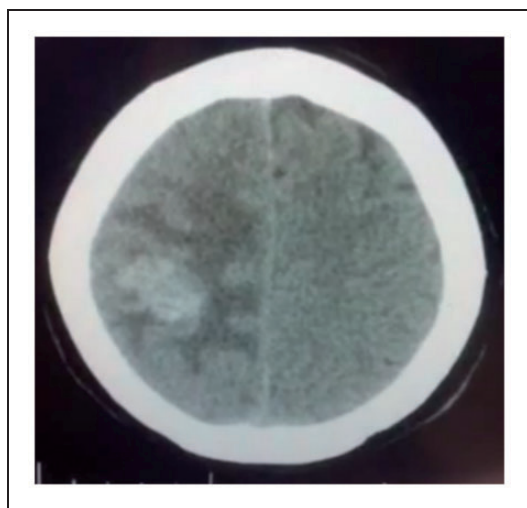


Figure 1. The axial brain computed tomography scan shows a hyperdense mass with surrounding hypodensity.

Intracranial calcification can be physiological as occurs in the basal ganglia and cerebellum due to the rich source of endogenous iron and calcium,² or can be associated with the pathological process as a slowly progressive glioma; and in infectious diseases such as tuberculosis, fungal infection and cysticercosis; and occasionally, with cystic lesions due to the slowing process of calcium deposition.⁶ Although the exact mechanism of calcification into metastatic lesions is unknown, there are two theories: first, degenerative and necrotic tissue decreases metabolism, resulting in disruption of carbon dioxide production, ultimately making the lesion alkaline compared to the surrounding parenchyma, consequently facilitating the deposition of calcium in alkaline tissue. Second, necrotic and degenerative tissue has a low level of oxygen that results in the influx of calcium into the intracellular space and a high level of alkaline phosphatase resulting in calcium phosphate deposition.²

Because calcification usually occurs in the necrotic part of the lesion, significant enhancement does not occur in this part so the peripheral non-necrotic part only shows marked enhancement.⁵

The most common intracranial masses with calcification includes oligodendroglioma, low-grade astrocytoma, craniopharyngioma, meningioma, pineal gland tumours and ependimoma.⁶

Although lung, breast, colon and ovaries are the most common primary site of cancer resulting in calcified brain metastasis, however, cases have been reported of calcified brain metastasis with an origin of squamous cell carcinoma and adenocarcinoma of the lung, sarcoma of the mediastinum, squamous cell carcinoma of the cervix, adenocarcinoma of the pancreas and non-Hodgkin's lymphoma.²

In the present case, the brain lesion was a solitary mass, hyperdense on CT scan associated with vasogenic oedema. Intralésional haemorrhage was the most differential diagnosis versus calcification. With the help of

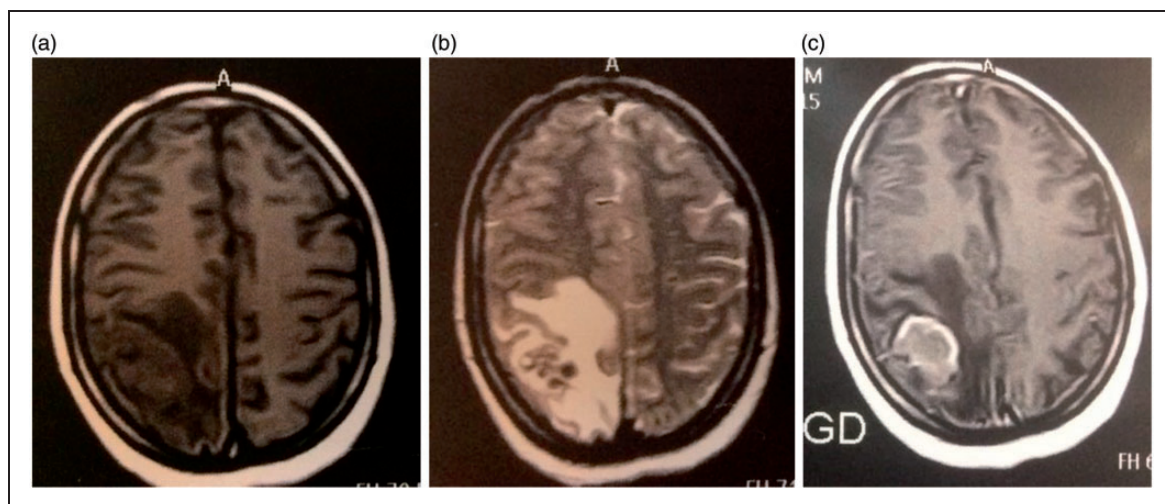


Figure 2. (a) Axial T1; (b) axial T2; and (c) T1 with gadolinium demonstrates the intra-axial lesion.

Table 1. Previous studies reporting cases of calcified brain metastasis.

| Row | Study | Age | Sex | Previous malignancy | Features of calcification | Location | Number of lesions | Procedure | Outcome/ follow-up duration | Confirm after | Histopathology |
|-----|---------------------------------|-----|-----|--|---|-----------------------|-------------------|-----------|-----------------------------|----------------|--|
| 1 | Callizo et al. ⁷ | 61 | M | - | Punctate/curvilinear comma-like scattered | Disseminate | Multiple | - | - | Autopsy | Acinar cell carcinoma of the pancreas |
| 2 | Ricke et al. ¹ | 52 | F | Ovarian cancer | Circular/ irregular | Disseminate | Multiple | Biopsy | - | Histopathology | Papillary carcinoma of the ovaries |
| 3 | Fukuda et al. ⁸ | 60 | F | Lung adenocarcinoma | Nodular | Disseminate | Multiple | - | - | Autopsy | Papillary adenocarcinoma lung |
| 4 | Gaze et al. ⁹ | 32 | w | Squamous carcinoma of cervix | Punctate | Temporoparietal | Solitary | Surgery | Death/10 months | Histopathology | Squamous carcinoma of cervix |
| 5 | Yamada and Suzuki ¹⁰ | 55 | M | Diffuse, large cell non-Hodgkin's lymphoma | Nodular | Thalamic region | Solitary | - | Survived/12 months | - | - |
| 6 | Tashiro et al. ³ | 57 | F | - | Conglomerate | Corpus callosum | Solitary | Surgery | Death/12 months | Histopathology | Metastatic squamous cell carcinoma/ unknown origin |
| 7 | Tomita and Larsen ¹¹ | 4.5 | M | Undifferentiated sarcoma | - | Parietal/ cerebellum | Two lesions | Surgery | ~3 Months | Histopathology | Metastatic undifferentiated sarcoma |
| 8 | Yamazaki et al. ¹² | 58 | M | - | Scattered punctate | Disseminate | Multiple | Biopsy | Death/8 months | Histopathology | Metastatic adenocarcinoma of the lung |
| 9 | Ohmoto et al. ¹³ | 29 | M | - | - | Frontal lobe | Solitary | Surgery | Survived/2 years & 3 months | Histopathology | Adenocarcinoma of lung |
| 10 | Kawamura et al. ² | 46 | F | Lung cancer | Linear | Frontal lobe | Solitary | Surgery | Death/3 years & 3 months | Histopathology | Adenocarcinoma |
| 11 | Bhatore and Gill ¹⁴ | 33 | F | Ovarian cancer | - | Cerebellar hemisphere | Solitary | Surgery | Death/12 days | Histopathology | Serous mucinous adenocarcinoma |
| 12 | Inomata et al. ¹⁵ | 68 | F | Lung adenocarcinoma | Scattered | Disseminate | Multiple | - | Death/1 year | Autopsy | Lung adenocarcinoma/ leptomeningeal adenocarcinoma |
| 13 | Fatehi et al. ¹⁶ | 30 | M | Breast cancer | - | Disseminate | Multiple | - | - | - | - |
| 14 | Michail et al. ¹⁷ | 60 | F | Colorectal carcinoma | - | Frontoparietal | Solitary | Surgery | - | Histopathology | Adenocarcinoma of colorectal |
| 15 | Stadnik et al. ¹⁸ | 70 | F | - | Peripheral calcification | Post fossa | Multiple | Surgery | - | Histopathology | Papillary cystadenocarcinoma |
| 16 | Nakase et al. ¹⁹ | 62 | F | Small cell carcinoma of lung | - | Disseminate | Multiple | - | Death/1 year | Autopsy | Small cell carcinoma of lung |

(continued)

Table 1. Continued

| Row | Study | Age | Sex | Previous malignancy | Features of calcification | Location | Number of lesions | Procedure | Outcome/follow-up duration | Confirm after | Histopathology |
|-----|--------------------------------|----------|---------|-------------------------------------|---------------------------|-----------------------------------|-------------------|-----------|----------------------------|----------------|--|
| 17 | Eom and Kim ⁶ | 50 | F | Lung cancer | - | Frontal lobe | solitary | Surgery | - | Histopathology | Non-small cell carcinoma of lung |
| 18 | Hwang et al. ²⁰ | 62 | M | - | - | Parietal/temporal/corpus callosum | Three lesions | Surgery | Death/2 years & 9 months | Histopathology | Metastatic adenocarcinoma |
| | | 50 | M | Squamous cell carcinoma of the lung | - | Temporal lobe | Solitary | Surgery | Death/1 year | Histopathology | Squamous cell carcinoma of the lung |
| 19 | Teksam et al. ²¹ | 21 | M | Osteosarcoma | - | Temporoparieto-occipital | Solitary | Surgery | Death/20 days | Histopathology | Calcified metastasis of osteosarcoma |
| 20 | Anand and Potts ⁵ | 38 to 72 | 5F/ 2 M | Lung, breast | Punctate | Temporal lobe | Solitary | - | - | - | - |
| | | | | Lung | Punctate | Supra/intra-tentorial | Multiple | - | - | - | - |
| | | | | Oesophagus | Curvilinear | Cerebellum | Solitary | - | - | - | - |
| | | | | Lung | Curvilinear | Cerebellum | Solitary | - | - | - | - |
| | | | | Colon | Amorphous | Frontal lobe | Solitary | - | - | - | - |
| | | | | Breast | Punctate | Supratentorial | Multiple | - | - | - | - |
| | | | | Colon | Amorphous | Frontoparietal | Solitary | - | - | - | - |
| 21 | Fernandez et al. ²² | 48 | M | Osteosarcoma of mandible | Round | Centrum semiovale | Two | - | - | Death/few days | - |
| 22 | Our study | 44 | F | - | Amorphous | Parietal lobe | Solitary | Surgery | Under follow-up | Histopathology | Adenocarcinoma, probably from colon origin |

Hounsfield units on CT scan (60 Hounsfield units) and MRI findings, the first diagnosis was a metastatic calcified mass confirmed intraoperatively. Considering the intraoperative observation, intraparenchymal meningioma was the most important differential diagnosis (due to shape, consistency and calcification of tumour). Ultimately, histopathological examination confirmed the diagnosis of metastatic adenocarcinoma. A thorough follow-up demonstrated that the caecum of the large intestine was the origin site of the tumour without additional dissemination of the primary lesion except for brain involvement. A retrograde history demonstrated that the patient had no gastrointestinal symptoms, and the brain lesion-related symptoms were the presenting symptoms of her cancer. After a 4-month follow-up, the patient was neurologically stable with no brain recurrence, while she was undergoing systemic chemotherapy. Table 1 demonstrates the most important studies, reporting cases of calcified brain metastasis.

Conclusion

Metastatic brain lesion should be in the differential diagnosis of a solitary calcified brain mass, although it occurs rarely. It is important to differentiate it radiologically from an intralesional haemorrhage.

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Conflict of interest

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